Package ‘FRASER’

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Type  Package
Title  Find RAre Splicing Events in RNA-Seq Data
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Description  Detection of rare aberrant splicing events in transcriptome profiles. Read count ratio expectations are modeled by an autoencoder to control for confounding factors in the data. Given these expectations, the ratios are assumed to follow a beta-binomial distribution with a junction specific dispersion. Outlier events are then identified as read-count ratios that deviate significantly from this distribution. FRASER is able to detect alternative splicing, but also intron retention. The package aims to support diagnostics in the field of rare diseases where RNA-seq is performed to identify aberrant splicing defects.

biocViews  RNASeq, AlternativeSplicing, Sequencing, Software, Genetics, Coverage

License  MIT + file LICENSE

URL  https://github.com/gagneurlab/FRASER

BugReports  https://github.com/gagneurlab/FRASER/issues

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Suggests  magick, BiocStyle, knitr, rmarkdown, testthat, covr,TxDb.Hsapiens.UCSC.hg19.knownGene, org.Hs.eg.db,
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annotateRanges

Annotates the given FraserDataSet with the HGNC symbol with biomaRt

Description

Annotates the given FraserDataSet with the HGNC symbol with biomaRt

Usage

```r
annotateRanges(
  fds,
  feature = "hgnc_symbol",
  featureName = feature,
  biotype = list("protein_coding"),
  ensembl = NULL,
  GRCh = 37
)
```

```r
annotateRangesWithTxDb(
  fds,
  feature = "SYMBOL",
  featureName = "hgnc_symbol",
  keytype = "ENTREZID",
  txdb = NULL,
  orgDb = NULL
)
```

Arguments

- **fds**: FraserDataSet
- **feature**: Defines which feature (default is HGNC symbol) should be annotated. Has to be the biomaRt feature name or a column name in orgDb.
- **featureName**: The column name of the feature in the FraserDataSet mcols.
- **biotype**: The biotype for biomaRt.
- **ensembl**: The ensembl that should be used. If NULL, the default one is used (hsapiens_gene_ensembl, GRCh37).
- **GRCh**: GRCh version to connect to. If this is NULL, then the current GRCh38 is used. Otherwise, this can only be 37 (default) at the moment (see useEnsembl).
The keytype or column name of gene IDs in the TxDb object (see keytypes for a list of available ID types).

A TxDb object. If this is NULL, then the default one is used, currently this is TxDb.Hsapiens.UCSC.hg19.knownGene.

An orgDb object or a data table to map the feature names. If this is NULL, then org.Hs.eg.db is used as the default.

Value

FraserDataSet

Examples

fds <- createTestFraserDataSet()

### Two ways to annotate ranges with gene names:
# either using biomart with GRCh38
try({
  fds <- annotateRanges(fds, GRCh=38)
  rowRanges(fds, type="psi5")[,c("hgnc_symbol")]
})

# either using biomart with GRCh37
try({
  fds <- annotateRanges(fds, featureName="hgnc_symbol_37", GRCh=37)
  rowRanges(fds, type="psi5")[,c("hgnc_symbol_37")]
})

# or with a provided TxDb object
require(TxDb.Hsapiens.UCSC.hg19.knownGene)
txdb <- TxDb.Hsapiens.UCSC.hg19.knownGene
require(org.Hs.eg.db)
orgDb <- org.Hs.eg.db
fds <- annotateRangesWithTxDb(fds, txdb=txdb, orgDb=orgDb)
rowRanges(fds, type="psi5")[,"hgnc_symbol"]

assayNames,FraserDataSet-method

Returns the assayNames of FRASER

Description

Returns the assayNames of FRASER

Usage

## S4 method for signature 'FraserDataSet'
assayNames(x)
assays,FraserDataSet-method

Arguments
x FraserDataSet

Value
Character vector

Description
Returns the assay for the given name/index of the FraserDataSet

Usage
## S4 method for signature 'FraserDataSet'
assays(x, withDimnames = TRUE, ...)

## S4 replacement method for signature 'FraserDataSet,SimpleList'
assays(x, withDimnames = TRUE, HDF5 = TRUE, type = NULL, ...) <- value

## S4 replacement method for signature 'FraserDataSet,list'
assays(x, withDimnames = TRUE, HDF5 = TRUE, type = NULL, ...) <- value

## S4 replacement method for signature 'FraserDataSet,DelayedMatrix'
assays(x, withDimnames = TRUE, HDF5 = TRUE, type = NULL, ...) <- value

Arguments
x FraserDataSet
withDimnames Passed on to SummarizedExperiment::assays()
... Parameters passed on to SummarizedExperiment::assays()
HDF5 Logical value indicating whether the assay should be stored as a HDF5 file.
type The psi type.
value The new value to which the assay should be set.

Value
(Delayed) matrix.
**calculatePSIValues**

*PSI value calculation*

**Description**

This function calculates the PSI values for each junction and splice site based on the FraserDataSet object.

**Usage**

```r
calculatePSIValues(
  fds,
  types = psiTypes,
  overwriteCts = FALSE,
  BPPARAM = bpparam()
)
```

**Arguments**

- `fds`  
  A *FraserDataSet* object

- `types`  
  A vector with the psi types which should be calculated. Default is all of psi5, psi3 and theta.

- `overwriteCts`  
  FALSE or TRUE (the default) the total counts (aka N) will be recalculated based on the existing junction counts (aka K)

- `BPPARAM`  
  the BiocParallel parameters for the parallelization

**Value**

*FraserDataSet*

**Examples**

```r
fds <- createTestFraserDataSet()
fds <- calculatePSIValues(fds, types="psi5")

### usually one would run this function for all psi types by using:
# fds <- calculatePSIValues(fds)
```
countRNA

Count RNA-seq data

Description

The FRASER package provides multiple functions to extract and count both split and non-spliced reads from bam files. See Detail and Functions for more information.

Usage

countRNAData(
  fds,
  NcpuPerSample = 1,
  minAnchor = 5,
  recount = FALSE,
  BPPARAM = bpparam(),
  genome = NULL,
  junctionMap = NULL,
  filter = TRUE,
  minExpressionInOneSample = 20,
  keepNonStandardChromosomes = TRUE,
  countDir = file.path(workingDir(fds), "savedObjects", nameNoSpace(name(fds))),
  ...
)

getSplitReadCountsForAllSamples(
  fds,
  NcpuPerSample = 1,
  junctionMap = NULL,
  recount = FALSE,
  BPPARAM = bpparam(),
  genome = NULL,
  countFiles = NULL,
  keepNonStandardChromosomes = TRUE,
  outDir = file.path(workingDir(fds), "savedObjects", nameNoSpace(name(fds)),
                  "splitCounts")
)

getNonSplitReadCountsForAllSamples(
  fds,
  splitCountRanges,
  NcpuPerSample = 1,
  minAnchor = 5,
  recount = FALSE,
  BPPARAM = bpparam(),
  longRead = FALSE,
  outDir = file.path(workingDir(fds), "savedObjects", nameNoSpace(name(fds)),
                    "nonSplitCounts")
)
addCountsToFraserDataSet(fds, splitCounts, nonSplitCounts)

countSplitReads(
    sampleID,
    fds,
    NcpuPerSample = 1,
    genome = NULL,
    recount = FALSE,
    keepNonStandardChromosomes = TRUE,
    bamfile = bamFile(fds[, sampleID]),
    pairedend = pairedEnd(fds[, sampleID]),
    strandmode = strandSpecific(fds),
    cacheFile = getSplitCountCacheFile(sampleID, fds),
    scanbamparam = scanBamParam(fds),
    coldata = colData(fds)
)

mergeCounts(
    countList,
    fds,
    junctionMap = NULL,
    assumeEqual = FALSE,
    spliceSiteCoords = NULL,
    BPPARAM = SerialParam()
)

countNonSplicedReads(
    sampleID,
    splitCountRanges,
    fds,
    NcpuPerSample = 1,
    minAnchor = 5,
    recount = FALSE,
    spliceSiteCoords = NULL,
    longRead = FALSE
)

**Arguments**

- **fds** A FraserDataSet object
- **NcpuPerSample** A BiocParallel param object or a positive integer to configure the parallel backend of the internal loop per sample
- **minAnchor** Minimum overlap around the Donor/Acceptor for non spliced reads. Default to 5
- **recount** if TRUE the cache is ignored and the bam file is recounted.
BPPARAM the BiocParallel parameters for the parallelization

genome NULL (default) or a character vector specifying the names of the reference genomes that were used to align the reads for each sample. The names have to be in a way accepted by the getBSgenome function. Available genomes can be listed using the available.genomes function from the BSgenome package. If genome is of length 1, the same reference genome will be used for all samples. If genome is supplied and strandSpecific(fds) == 0L (unstranded), then the strand information will be estimated by checking the dinucleotides found at the intron boundaries (see summarizeJunctions in GenomicAlignments package for details). This can e.g. help to avoid ambiguities when adding gene names from a gene annotation to the introns in a later step.

junctionMap A object or file containing a map of all junctions of interest across all samples

filter If TRUE, splice sites of introns with low read support in all samples are not considered when calculating the non-split reads. This helps to speed up the subsequent steps.

minExpressionInOneSample The minimal split read count in at least one sample that is required for an intron to pass the filter.

keepNonStandardChromosomes Logical value indicating if non standard chromosomes should also be counted. Defaults to TRUE.

countDir The directory in which the tsv containing the position and counts of the junctions should be placed.

... Further parameters passed on to Rsubread::featureCounts.

countFiles If specified, the split read counts for all samples are read from the specified files. Should be a vector of paths to files containing the split read counts for the individual samples. Reading from files is only supported for tsv(.gz) or RDS files containing GRranges objects. The order of the individual sample files should correspond to the order of the samples in the fds.

outDir The full path to the output folder containing the merged counts. If the given folder already exists and stores a SummarizedExperiment object, the counts from this folder will be read in and used in the following (i.e. the reads are not recounted), unless the option recount=TRUE is used. If this folder doesn’t exist or if recount=TRUE, then it will be created after counting has finished.

splitCountRanges The merged GRanges object containing the positions of all the introns in the dataset over all samples.

longRead If TRUE, then the isLongRead option of Rsubread::featureCounts is used when counting the non spliced reads overlapping splice sites.

splitCounts The SummarizedExperiment object containing the position and counts of all the introns in the dataset for all samples.

nonSplitCounts The SummarizedExperiment object containing the position and non split read counts of all splice sites present in the dataset for all samples.

currentID The ID of the sample to be counted.
bamfile

The BAM file to be used to extract the counts. Defaults to the BAM file defined in the FraserDataSet object.

pairedend

TRUE or FALSE if the BAM file is paired end. Defaults to the value specified in the FraserDataSet object.

strandmode

0 (no, default), 1 (stranded), or 2 (revers) to specify the used protocol for the RNA-seq experiment.

cacheFile

File path to the cache, where counts are stored.

scanbamparam

The ScanBamParam object which is used for loading the reads from the BAM file before counting. Defaults to the params stored in the FraserDataSet object.

coldata

The colData as given by the FraserDataSet object.

countList

A list of GRanges objects containing the counts that should be merged into one object.

assumeEqual

Logical indicating whether all objects in countList can be assumed to contain counts for the same ranges. If FALSE, merging of the ranges is performed.

spliceSiteCoords

A GRanges object containing the positions of the splice sites. If it is NULL, then splice sites coordinates are calculated first based on the positions of the junctions defined from the split reads.

Details

The functions described in this file extract and count both the split and the non-spliced reads from bam files.

countRNAData is the main function that takes care of all counting steps and returns a FraserDataSet containing the counts for all samples in the fds.

getSplitReadCountsForAllSamples counts split reads for all samples and getNonSplitReadCountsForAllSamples counts non split reads overlapping splice sites for all samples. addCountsToFraserDataSet adds these counts to an existing fds.

countSplitReads calculates the split read counts for a single sample. countNonSplicedReads counts the non split reads overlapping with splice sites for a single sample.

mergeCounts merges the counts from different samples into a single count object, where the counts for junctions that are not present in a sample are set to zero.

Value

countRNAData returns a FraserDataSet.
getSplitReadCountsForAllSamples returns a GRanges object.
getNonSplitReadCountsForAllSamples returns a GRanges object.
addCountsToFraserDataSet returns a FraserDataSet.
countSplitReads returns a GRanges object.
mergeCounts returns a SummarizedExperiment object.
countNonSplicedReads returns a GRanges object.
createTestFraserSettings

Functions

- `countRNAData()`: This method extracts and counts the split reads and non spliced reads from RNA bam files.
- `getSplitReadCountsForAllSamples()`: This method creates a GRanges object containing the split read counts from all specified samples.
- `getNonSplitReadCountsForAllSamples()`: This method creates a GRanges object containing the non split read counts at the exon-intron boundaries inferred from the GRanges object containing the positions of all the introns in this dataset.
- `addCountsToFraserDataSet()`: This method adds the split read and non split read counts to a existing FraserDataSet containing the settings.
- `countSplitReads()`: This method counts all split reads in a bam file for a single sample.
- `mergeCounts()`: This method merges counts for multiple samples into one SummarizedExperiment object.
- `countNonSplicedReads()`: This method counts non spliced reads based on the given target (acceptor/donor) regions for a single sample.

Examples

```r
# On Windows SNOW is the default for the parallel backend, which can be
# very slow for many but small tasks. Therefore, we will use
# for the example the SerialParam() backend.
if(!.Platform$OS.type == "unix") {
  register(SerialParam())
}

fds <- countRNAData(createTestFraserSettings())
```

createTestFraserSettings

Create a test dataset

Description

Create a test case dataset based on the test sample annotation to be used in the vignette and to explore the functionality of the FRASER package. Dependent on the request only the sample annotation or a full fitted model is returned.

Usage

```r
createTestFraserSettings(workingDir = "FRASER_output")
createTestFraserDataSet(workingDir = "FRASER_output", rerun = FALSE)
```
filtering

Arguments

- **workingDir**: Directory where to store HDF5 and RDS files. Defaults to `FRASER_output` in the current working directory.
- **rerun**: Defaults to `FALSE`. If set to `TRUE` it reruns the full fit of the model.

Value

A FraserDataSet object that contains a test case

Examples

```r
fds <- createTestFraserSettings()
fds

fds <- createTestFraserDataSet()
fds
```

filtering  Filtering FraserDataSets

Description

This method can be used to filter out introns that are not reliably detected and to remove introns with no variability between samples.

Usage

```r
filterExpressionAndVariability(
  object,
  minExpressionInOneSample = 20,
  quantile = 0.95,
  quantileMinExpression = 10,
  minDeltaPsi = 0.05,
  filter = TRUE,
  delayed = ifelse(ncol(object) <= 300, FALSE, TRUE),
  BPPARAM = bpparam()
)
```

```r
## S4 method for signature 'FraserDataSet'
filterExpression(
  object,
  minExpressionInOneSample = 20,
  quantile = 0.95,
  quantileMinExpression = 10,
  filter = TRUE,
  delayed = ifelse(ncol(object) <= 300, FALSE, TRUE),
  BPPARAM = bpparam()
)
```
filtering

BPPARAM = bpparam()

filterVariability(
  object,
  minDeltaPsi = 0.05,
  filter = TRUE,
  delayed = ifelse(ncol(object) <= 300, FALSE, TRUE),
  BPPARAM = bpparam()
)

Arguments

object A FraserDataSet object

minExpressionInOneSample The minimal read count in at least one sample that is required for an intron to pass the filter.

quantile Defines which quantile should be considered for the filter.

quantileMinExpression The minimum read count an intron needs to have at the specified quantile to pass the filter.

minDeltaPsi Only introns for which the maximal difference in the psi value of a sample to the mean psi of the intron is larger than this value pass the filter.

filter If TRUE, a subsetted fds containing only the introns that passed all filters is returned. If FALSE, no subsetting is done and the information of whether an intron passed the filters is only stored in the mcols.

delayed If FALSE, count matrices will be loaded into memory, otherwise the function works on the delayedMatrix representations. The default value depends on the number of samples in the fds-object.

BPPARAM the BiocParallel parameters for the parallelization

Value

A FraserDataSet with information about which junctions passed the filters. If filter=TRUE, the filtered FraserDataSet is returned.

Functions

- filterExpressionAndVariability(): This functions filters out both introns with low read support and introns that are not variable across samples.
- filterExpression(FraserDataSet): This function filters out introns and corresponding splice sites that have low read support in all samples.
- filterVariability(): This function filters out introns and corresponding splice sites which do not show variability across samples.
Examples

```r
fds <- createTestFraserDataSet()
# filterExpressionAndVariability(fds, minDeltaPsi=0.1, filter=FALSE)
mcols(fds, type="psi5")[, c(
  "maxCount", "passedExpression", "maxDPsi3", "passedVariability")]
plotFilterExpression(fds)
plotFilterVariability(fds)
```

---

**fit**

*Fitting the denoising autoencoder*

---

**Description**

This method corrects for confounders in the data and fits a beta-binomial distribution to the introns/splice sites.

For more details please see [FRASER](https://github.com/NestorVila/FRASER).

**Usage**

```r
## S3 method for class 'FraserDataSet'
fit(
  object,
  implementation = c("PCA", "PCA-BB-Decoder", "AE", "AE-weighted", "PCA-BB-full",
                     "fullAE", "PCA-regression", "PCA-reg-full", "PCA-BB-Decoder-no-weights", "BB"),
  q,
  type = "psi3",
  rhoRange = c(1e-08, 1 - 1e-08),
  weighted = FALSE,
  noiseAlpha = 1,
  convergence = 1e-05,
  iterations = 15,
  initialize = TRUE,
  control = list(),
  BPPARAM = bpparam(),
  nSubset = 15000,
  minDeltaPsi = 0.1,
  ...
)
```

**Arguments**

- `object`  
  A `FraserDataSet` object

- `implementation`  
  The method that should be used to correct for confounders.
The encoding dimensions to be used during the fitting procedure. Should be fitted using `optimHyperParams` if unknown. If a named vector is provided it is used for the different splicing types.

**type**
The type of PSI (psi5, psi3 or theta for theta/splicing efficiency)

**rhoRange**
Defines the range of values that rho parameter from the beta-binomial distribution is allowed to take. For very small values of rho, the loss can be instable, so it is not recommended to allow rho < 1e-8.

**weighted**
If TRUE, the weighted implementation of the autoencoder is used

**noiseAlpha**
Controls the amount of noise that is added for the denoising autoencoder.

**convergence**
The fit is considered to have converged if the difference between the previous and the current loss is smaller than this threshold.

**iterations**
The maximal number of iterations. When the autoencoder has not yet converged after these number of iterations, the fit stops anyway.

**initialize**
If FALSE and a fit has been previously run, the values from the previous fit will be used as initial values. If TRUE, (re-)initialization will be done.

**control**
List of control parameters passed on to optim().

**BPPARAM**
the BiocParallel parameters for the parallelization

**nSubset**
The size of the subset to be used in fitting if subsetting is used.

**minDeltaPsi**
Minimal delta psi of an intron to be be considered a variable intron.

... Currently not used

### Value

**FraserDataSet**

### See Also

**FRASER**
Usage

FRASER(
  fds,
  q,
  implementation = c("PCA", "PCA-BB-Decoder", "AE-weighted", "AE", "BB"),
  iterations = 15,
  BPPARAM = bpparam(),
  correction,
  ...
)

calculateZscore(fds, type = currentType(fds), logit = TRUE)

calculatePvalues(
  fds,
  type = currentType(fds),
  implementation = "PCA",
  BPPARAM = bpparam(),
  distributions = c("betabinomial"),
  capN = 5 * 1e+05
)

calculatePadjValues(fds, type = currentType(fds), method = "BY")

Arguments

fds A FraserDataSet object
q The encoding dimensions to be used during the fitting procedure. Should be
    fitted using optimHyperParams if unknown. If a named vector is provided it is
    used for the different splicing types.
implementation The method that should be used to correct for confounders.
iterations The maximal number of iterations. When the autoencoder has not yet converged
    after these number of iterations, the fit stops anyway.
BPPARAM A BiocParallel object to run the computation in parallel
correction Deprecated. The name changed to implementation.
... Additional parameters passed on to the internal fit function
type The type of PSI (psi5, psi3 or theta for theta/splicing efficiency)
logit Indicates if z scores are computed on the logit scale (default) or in the natural
    (psi) scale.
distributions The distribution based on which the p-values are calculated. Possible are beta-
    binomial, binomial and normal.
capN Counts are capped at this value to speed up the p-value calculation
method The p.adjust method that should be used.
Details

All computed values are returned as an FraserDataSet object. To have more control over each analysis step, one can call each function separately.

- `fit` to control for confounding effects and fit the beta binomial model parameters
- `calculatePvalues` to calculate the nominal p values
- `calculatePadjValues` to calculate adjusted p values (per sample)
- `calculateZscore` to calculate the Z scores

Available methods to correct for the confounders are currently: a denoising autoencoder with a BB loss ("AE" and "AE-weighted"), PCA ("PCA"), a hybrid approach where PCA is used to fit the latent space and then the decoder of the autoencoder is fit using the BB loss ("PCA-BB-Decoder"). Although not recommended, it is also possible to directly fit the BB distribution to the raw counts ("BB").

Value

FraserDataSet

Functions

- `FRASER()`: This function runs the default FRASER pipeline combining the beta-binomial fit, the computation of Z scores and p values as well as the computation of delta-PSI values.
- `calculateZscore()`: This function calculates z-scores based on the observed and expected logit psi.
- `calculatePvalues()`: This function calculates two-sided p-values based on the beta-binomial distribution (or binomial or normal if desired). The returned p values are already adjusted with Holm's method per donor or acceptor site, respectively.
- `calculatePadjValues()`: This function adjusts the previously calculated p-values per sample for multiple testing.

Author(s)

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See Also

`fit`

Examples

```r
# set default parallel backend
register(SerialParam())

# preprocessing
fds <- createTestFraserDataSet()
```
# filtering not expressed introns
fds <- calculatePSIValues(fds)
fds <- filterExpressionAndVariability(fds)

# Run the full analysis pipeline: fits distribution and computes p values
fds <- FRASER(fds, q=2, implementation="PCA")

# afterwards, the fitted fds-object can be saved and results can
# be extracted and visualized, see ?saveFraserDataSet, ?results and
# ?plotVolcano

# The functions run inside the FRASER function can also be directly
# run themselves.
# To directly run the fit function:
fds <- fit(fds, implementation="PCA", q=2, type="psi5")

# To directly run the nomial and adjusted p value and z score
# calculation, the following functions can be used:
fds <- calculatePvalues(fds, type="psi5")
head(pVals(fds, type="psi5"))
fds <- calculatePadjValues(fds, type="psi5", method="BY")
head(padjVals(fds, type="psi5"))
fds <- calculateZscore(fds, type="psi5")
head(zScores(fds, type="psi5"))

---

**FraserDataSet**  
*The FRASER dataset object*

**Description**

Constructs an FRASER object based on the given input. It can take only the annotation (colData) or count tables (junctions/spliceSites).

**Usage**

FraserDataSet(colData = NULL, junctions = NULL, spliceSites = NULL, ...)

**Arguments**

- **colData**: A DataFrame containing the annotation of the samples
- **junctions, spliceSites**: A data.frame like containing the raw counts for each junction or splice site. It requires the columns startID and endID for the junctions and spliceSiteID and type for the splice sites. Those columns identifies the corresponding splice site for the given junction and map to the splice site. For each sample the counts are saved in a corresponding column with the same name. It can also be a GRRange object.
- **...**: Any parameters corresponding to the slots and their possible values. See FraserDataSet
FraserDataSet-class

Value
A FraserDataSet object.

Author(s)
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Examples
fraser <- FraserDataSet()

# example sample annotation
sampleTable <- fread(system.file("extdata", "sampleTable_countTable.tsv", package="FRASER", mustWork=TRUE))

# get raw counts
junctionCts <- fread(system.file("extdata", "raw_junction_counts.tsv.gz", package="FRASER", mustWork=TRUE))
spliceSiteCts <- fread(system.file("extdata", "raw_site_counts.tsv.gz", package="FRASER", mustWork=TRUE))

# create FRASER object
fds <- FraserDataSet(colData=sampleTable, junctions=junctionCts, spliceSites=spliceSiteCts, name="Example Dataset")

---

FraserDataSet-class  FraserDataSet

Description
This class is designed to store the whole FRASER data set needed for an analysis of a disease cohort

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---

getter_setter_functions

Getter/Setter functions

Description
This is a collection of small accessor/setter functions for easy access to the values within the FRASER model.
Usage

featureExclusionMask(fds, type = currentType(fds))

rho(fds, type = currentType(fds))

zScores(fds, type = currentType(fds), byGroup = FALSE, ...)

pVals(fds, type = currentType(fds), level = "site", dist = "BetaBinomial", ...)

padjVals(fds, type = currentType(fds), dist = c("BetaBinomial"), ...)

predictedMeans(fds, type = currentType(fds))

deltaPsiValue(fds, type = currentType(fds))

currentType(fds)

currentType(fds) <- value

pseudocount(value = NULL)

hyperParams(fds, type = currentType(fds), all = FALSE)

bestQ(fds, type = currentType(fds))

dontWriteHDF5(fds)

dontWriteHDF5(fds) <- value

verbose(fds)

verbose(fds) <- value

Arguments

fds An FraserDataSet object.
type The type of psi (psi5, psi3 or theta)
value The new value to be assigned.
byGroup If TRUE, aggregation by donor/acceptor site will be done.
... Internally used parameters.
level Indicates if the retrieved p values should be adjusted on the donor/acceptor site-level (default) or if unadjusted junction-level p values should be returned.
dist Distribution for which the p-values should be extracted.
Logical value indicating whether `hyperParams(fds)` should return the results of all evaluated parameter combinations or only for the optimal parameter combination.

**Value**

A (delayed) matrix or vector dependent on the type of data retrieved.

**Functions**

- `featureExclusionMask()`: Retrieves a logical vector indicating for each junction whether it is included or excluded during the fitting procedure.
- `featureExclusionMask(fds, type = currentType(fds)) <- value`: To remove certain junctions from being used in the train step of the encoding dimension we can set the `featureExclusion` vector to FALSE. This can be helpful if we have local linkage between features which we do not want to model by the autoencoder.
- `rho()`: Returns the fitted rho values for the beta-binomial distribution
- `zScores()`: This returns the calculated z-scores.
- `pVals()`: This returns the calculated p-values.
- `padjVals()`: This returns the adjusted p-values.
- `predictedMeans()`: This returns the fitted mu (i.e. psi) values.
- `deltaPsiValue()`: Returns the difference between the observed and the fitted psi values.
- `currentType()`: Returns the psi type that is used within several methods in the FRASER package.
- `currentType(fds) <- value`: Sets the psi type that is to be used within several methods in the FRASER package.
- `pseudocount()`: Sets and returns the pseudo count used within the FRASER fitting procedure.
- `hyperParams()`: This returns the results of the hyperparameter optimization NULL if the hyperparameter optimization was not run yet.
- `bestQ()`: This returns the optimal size of the latent space according to the hyperparameter optimization or a simple estimate of about a tenth of the number of samples if the hyperparameter optimization was not run yet.
- `dontWriteHDF5()`: Gets the current value of whether the assays should be stored as hdf5 files.
- `dontWriteHDF5(fds) <- value`: Sets whether the assays should be stored as hdf5 files.
- `verbose()`: Dependend on the level of verbosity the algorithm reports more or less to the user. 0 means being quiet and 10 means everything.
- `verbose(fds) <- value`: Sets the verbosity level to a value between 0 and 10. 0 means being quiet and 10 means reporting everything.
injectOutliers

Inject artificial outliers in an existing fds

Description

Inject artificial outliers in an existing fds

Usage

injectOutliers(
  fds,
  type = c("psi5", "psi3", "theta"),
  freq = 0.001,
  minDpsi = 0.2,
  minCoverage = 2,
)
deltaDistr = "uniformDistr",
verbose = FALSE,
method = c("samplePSI", "meanPSI", "simulatedPSI"),
BPPARAM = bpparam()
)

Arguments

fds FraserDataSet
type The psi type
dfreq The injection frequency.
minDpsi The minimal delta psi with which outliers will be injected.
minCoverage The minimal total coverage (i.e. N) required for a junction to be considered for injection of an outlier.
deltaDistr The distribution from which the delta psi value of the injections is drawn (default: uniform distribution).
verbose Should additional information be printed during computation?
method Defines by which method the new psi of injections is computed, i.e. to which value the delta psi of the injection is added: "meanPSI" for adding to the mean psi of the junction over all samples or "samplePSI" to add to the psi value of the junction in the specific sample. "simulatedPSI" is only possible if a simulated dataset is used.
BPPARAM A BiocParallel object to run the computation in parallel

Value

FraserDataSet

Examples

# A generic dataset
defs <- makeSimulatedFraserDataSet()
defs <- injectOutliers(defs, minDpsi=0.2, freq=1E-3)

K Getter/setter for count data

Description

Getter/setter for count data

setter for count data
Usage

\[ K(fds, \text{type} = \text{currentType}(fds)) \]

\[ N(fds, \text{type} = \text{currentType}(fds)) \]

## S4 method for signature 'FraserDataSet'

\[ \text{counts(object, type = NULL, side = c("ofInterest", "otherSide"))} \]

## S4 replacement method for signature 'FraserDataSet,ANY'

\[ \text{counts(object, type = NULL, side = c("ofInterest", "otherSide"), \ldots) <- value} \]

Arguments

- \text{fds, object} \quad \text{FraserDataSet}
- \text{type} \quad \text{The psi type.}
- \text{side} \quad \text{"ofInterest" for junction counts, "other" for sum of counts of all other junctions at the same donor site (psi5) or acceptor site (psi3), respectively.}
- \text{\ldots} \quad \text{Further parameters that are passed to assays(object,\ldots)}
- \text{value} \quad \text{An integer matrix containing the counts.}

Value

\text{FraserDataSet}

Examples

\begin{verbatim}
fds <- createTestFraserDataSet()

\text{counts(fds, type="psi5", side="ofInterest")}
\text{counts(fds, type="psi5", side="other")}
\text{head(K(fds, type="psi3"))}
\text{head(N(fds, type="theta"))}
\end{verbatim}

\hline
\text{length,FraserDataSet-method}

\begin{center}
\text{\textit{retrieve the length of the object (aka number of junctions)}}
\end{center}

Description

retrieve the length of the object (aka number of junctions)

Usage

## S4 method for signature 'FraserDataSet'

\[ \text{length(x)} \]
loadFraserDataSet

Arguments

x FraserDataSet

Value

Length of the object.

loadFraserDataSet Loading/Saving FraserDataSets

Description

This is a convenient function to load and save a FraserDataSet object. It looks and saves the FraserDataSet objects and HDF5 files on disk under the given working dir. Internally it uses HDF5 files for all assays.

Usage

loadFraserDataSet(dir, name = NULL, file = NULL, upgrade = FALSE)

saveFraserDataSet(fds, dir = NULL, name = NULL, rewrite = FALSE)

Arguments

dir A path where to save the objects (replaces the working directory)
name The analysis name of the project (saved within the 'dir')
file The file path to the fds-object.RDS file that should be loaded.
upgrade Should the version of the loaded object be updated?
fds A FraserDataSet object to be saved
rewrite logical if the object should be rewritten. This makes sense if you have filtered or subsetted the object and want to save only the subsetted version

Value

FraserDataSet

Examples

fds <- createTestFraserSettings()
name(fds) <- "saving_test"

# make sure the object is saved to disc
dontWriteHDF5(fds) <- FALSE
dfsSaved <- saveFraserDataSet(fds)
# fdsSaved
makeSimulatedFraserDataSet

Create an simulated example data set for FRASER

Description
Simulates a data set based on random counts following a beta binomial (or Dirichlet-Multinomial) distribution.

Usage
makeSimulatedFraserDataSet(
m = 100,
  j = 500,
  q = 10,
  distribution = c("BB", "DM"),
  ...
)

Arguments

m  Number of simulated samples
j  Number of simulated junctions
q  number of simulated latent variables.
distribution  Either "BB" for a beta-binomial simulation or "DM" for a dirichlet-multinomial simulation.
...  Further arguments used to construct the FraserDataSet.

Value
An FraserDataSet containing an example dataset based on simulated data

Examples

# A generic dataset
fds1 <- makeSimulatedFraserDataSet()
fd1

# A generic dataset with specified sample size and injection method
fds2 <- makeSimulatedFraserDataSet(m=10, j=100, q=3)
fds2
Description
To boost its own sequencing data, one can download existing and precounted data. This function merges the existing FraserDataSet with external count data.

Usage
mergeExternalData(fds, countFiles, sampleIDs, annotation = NULL)

Arguments
- **fds**
  A FraserDataSet
- **countFiles**
  A character vector of file names pointing to the external count data. The vector has to be names or the files have to start with \texttt{k\_j}, \texttt{k\_theta}, \texttt{n\_psi3}, \texttt{n\_psi5}, \texttt{n\_theta}.
- **sampleIDs**
  The samples to be merged from the external data.
- **annotation**
  A sample annotation of the external data (optional).

Details
For more details on existing datasets have a look at: \url{https://github.com/gagneurlab/drop#datasets}
Since FRASER can not hand NA values, the merge will return only the intersecting regions and will drop any non overlapping features. This has to be kept in mind when analysing rare disease samples.

Value
Merged FraserDataSet object.

Examples
```r
anno <- fread(system.file("extdata", "externalCounts", "annotation.tsv.gz", package="FRASER"))
cntsFiles <- list.files(full.names = TRUE, pattern="counts", system.file("extdata", "externalCounts", package="FRASER"))

data <- createTestFraserDataSet()
mergeExternalData(data, cntsFiles, anno[,sampleID], anno)
K(data, "psi5")
K(mergeExternalData(data, cntsFiles, anno[,sampleID], anno), "psi5")
```
optimHyperParams  

Find optimal encoding dimension

Description
---
Finds the optimal encoding dimension by injecting artificial splicing outlier ratios while maximizing
the precision-recall curve.

Usage
---
optimHyperParams(
  fds,
  type,
  implementation = "PCA",
  q_param = seq(2, min(40, ncol(fds)), by = 3),
  noise_param = 0,
  minDeltaPsi = 0.1,
  iterations = 5,
  setSubset = 50000,
  injectFreq = 0.01,
  BPPARAM = bpparam(),
  internalThreads = 1,
  plot = TRUE,
  delayed = ifelse(ncol(fds) <= 300, FALSE, TRUE),
  ...
)

Arguments
---
<table>
<thead>
<tr>
<th>Argument</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>fds</td>
<td>A FraserDataSet object</td>
</tr>
<tr>
<td>type</td>
<td>The type of PSI (psi5, psi3 or theta for theta/splicing efficiency)</td>
</tr>
<tr>
<td>implementation</td>
<td>The method that should be used to correct for confounders.</td>
</tr>
<tr>
<td>q_param</td>
<td>Vector specifying which values of q should be tested.</td>
</tr>
<tr>
<td>noise_param</td>
<td>Vector specifying which noise levels should be tested.</td>
</tr>
<tr>
<td>minDeltaPsi</td>
<td>Minimal delta psi of an intron to be considered a variable intron.</td>
</tr>
<tr>
<td>iterations</td>
<td>The maximal number of iterations. When the autoencoder has not yet converged</td>
</tr>
<tr>
<td></td>
<td>after these number of iterations, the fit stops anyway.</td>
</tr>
<tr>
<td>setSubset</td>
<td>The size of the subset of the most variable introns that should be used for</td>
</tr>
<tr>
<td></td>
<td>the hyperparameter optimization.</td>
</tr>
<tr>
<td>injectFreq</td>
<td>The frequency with which outliers are injected into the data.</td>
</tr>
<tr>
<td>BPPARAM</td>
<td>the BiocParallel parameters for the parallelization</td>
</tr>
<tr>
<td>internalThreads</td>
<td>The number of threads used internally.</td>
</tr>
</tbody>
</table>
plotFunctions

plot  If TRUE, a plot of the area under the curve and the model loss for each evaluated parameter combination will be displayed after the hyperparameter optimization finishes.

delayed  If FALSE, count matrices will be loaded into memory (faster calculations), otherwise the function works on the delayedMatrix representations (more memory efficient). The default value depends on the number of samples in the fds-object.

Additional parameters passed to injectOutliers.

Value  FraserDataSet

See Also  FRASER

Examples

# generate data
fds <- makeSimulatedFraserDataSet(m=15, j=20)

# run hyperparameter optimization
fds <- optimHyperParams(fds, type="psi5", q_param=c(2, 5))

# get estimated optimal dimension of the latent space
bestQ(fds, type="psi5")
hyperParams(fds, type="psi5")

plotFunctions  Visualization functions for FRASER

Description

The FRASER package provides multiple functions to visualize the data and the results of a full data set analysis.

Plots the p values over the delta psi values, known as volcano plot. Visualizes per sample the outliers. By type and aggregate by gene if requested.

Plot the number of aberrant events per samples

Plots the observed split reads of the junction of interest over all reads coming from the given donor/acceptor.

Plots the expected psi value over the observed psi value of the given junction.

Plots the quantile-quantile plot

Histogram of the geometric mean per junction based on the filter status

Histogram of minimal delta psi per junction

Count correlation heatmap function
Usage

## S4 method for signature 'FraserDataSet'
plotVolcano(
  object,
  sampleID,
  type = c("psi3", "psi5", "theta"),
  basePlot = TRUE,
  aggregate = FALSE,
  main = NULL,
  label = NULL,
  deltaPsiCutoff = 0.3,
  padjCutoff = 0.1,
  ...
)

## S4 method for signature 'FraserDataSet'
plotAberrantPerSample(
  object,
  main,
  type = c("psi3", "psi5", "theta"),
  padjCutoff = 0.1,
  zScoreCutoff = NA,
  deltaPsiCutoff = 0.3,
  aggregate = TRUE,
  BPPARAM = bpparam(),
  ...
)

plotExpression(
  fds,
  type = c("psi5", "psi3", "theta"),
  site = NULL,
  result = NULL,
  colGroup = NULL,
  basePlot = TRUE,
  main = NULL,
  label = "aberrant",
  ...
)

plotExpectedVsObservedPsi(
  fds,
  type = c("psi5", "psi3", "theta"),
  idx = NULL,
  result = NULL,
  colGroup = NULL,
  main = NULL,
  basePlot = TRUE,
label = "aberrant",
...
)

## S4 method for signature 'FraserDataSet'
plotQQ(
  object,
  type = NULL,
  idx = NULL,
  result = NULL,
  aggregate = FALSE,
  global = FALSE,
  main = NULL,
  conf.alpha = 0.05,
  samplingPrecision = 3,
  basePlot = TRUE,
  label = "aberrant",
  Ncpus = min(3, getDTthreads()),
  ...
)

## S4 method for signature 'FraserDataSet'
plotEncDimSearch(
  object,
  type = c("psi3", "psi5", "theta"),
  plotType = c("auc", "loss")
)

plotFilterExpression(
  fds,
  bins = 200,
  legend.position = c(0.8, 0.8),
  onlyVariableIntrons = FALSE
)

plotFilterVariability(
  fds,
  bins = 200,
  legend.position = c(0.8, 0.8),
  onlyExpressedIntrons = FALSE
)

## S4 method for signature 'FraserDataSet'
plotCountCorHeatmap(
  object,
  type = c("psi5", "psi3", "theta"),
  logit = FALSE,
  topN = 50000,
)
topJ = 5000,
minMedian = 1,
minCount = 10,
main = NULL,
normalized = FALSE,
show_rownames = FALSE,
show_colnames = FALSE,
minDeltaPsi = 0.1,
annotation_col = NA,
annotation_row = NA,
border_color = NA,
nClust = 5,
plotType = c("sampleCorrelation", "junctionSample"),
sampleClustering = NULL,
plotMeanPsi = TRUE,
plotCov = TRUE,
... )

Arguments

object, fds An FraserDataSet object.
sampleID A sample ID which should be plotted. Can also be a vector. Integers are treated as indices.
type The psi type: either psi5, psi3 or theta (for SE).
basePlot if TRUE (default), use the R base plot version, else use the plotly framework.
aggregate If TRUE, the pvalues are aggregated by gene (default), otherwise junction level pvalues are used (default for Q-Q plot).
main Title for the plot, if missing a default title will be used.
label Indicates the genes or samples that will be labelled in the plot (only for basePlot=TRUE).
Setting label="aberrant" will label all aberrant genes or samples. Labelling can be turned off by setting label=NULL. The user can also provide a custom list of gene symbols or sampleIDs.

padjCutoff, zScoreCutoff, deltaPsiCutoff Significance, Z-score or delta psi cutoff to mark outliers

... Additional parameters passed to plot() or plot_ly() if not stated otherwise in the details for each plot function

BPPARAM BiocParallel parameter to use.
result The result table to be used by the method.
colGroup Group of samples that should be colored.
idx, site A junction site ID or gene ID or one of both, which should be plotted. Can also be a vector. Integers are treated as indices.

conf.alpha If set, a confidence interval is plotted, defaults to 0.05
**plotFunctions**

- **samplingPrecision**: Plot only non-overlapping points in Q-Q plot to reduce number of points to plot. Defines the digits to round to.

- **Ncpus**: Number of cores to use.

- **plotType**: The type of plot that should be shown as character string. For plotEncDimSearch, it has to be either "auc" for a plot of the area under the curve (AUC) or "loss" for the model loss. For the correlation heatmap, it can be either "sampleCorrelation" for a sample-sample correlation heatmap or "junctionSample" for a junction-sample correlation heatmap.

- **bins**: Set the number of bins to be used in the histogram.

- **legend.position**: Set legend position (x and y coordinate), defaults to the top right corner.

- **onlyVariableIntrons**: Logical value indicating whether to show only introns that also pass the variability filter. Defaults to FALSE.

- **onlyExpressedIntrons**: Logical value indicating whether to show only introns that also pass the expression filter. Defaults to FALSE.

- **logit**: If TRUE, the default, psi values are plotted in logit space.

- **topN**: Top x most variable junctions that should be used for the calculation of sample x sample correlations.

- **topJ**: Top x most variable junctions that should be displayed in the junction-sample correlation heatmap. Only applies if plotType is "junctionSample".

- **minMedian, minCount, minDeltaPsi**: Minimal median ($m \geq 1$), delta psi ($|\Delta \psi| > 0.1$), read count ($n \geq 10$) value of a junction to be considered for the correlation heatmap.

- **normalized**: If TRUE, the normalized psi values are used, the default, otherwise the raw psi values

- **show_rownames, show_colnames**: Logical value indicating whether to show row or column names on the heatmap axes.

- **annotation_col, annotation_row**: Row or column annotations that should be plotted on the heatmap.

- **border_color**: Sets the border color of the heatmap.

- **nClust**: Number of clusters to show in the row and column dendrograms.

- **sampleClustering**: A clustering of the samples that should be used as an annotation of the heatmap.

- **plotMeanPsi, plotCov**: If TRUE, then the heatmap is annotated with the mean psi values or the junction coverage.

**Details**

This is the list of all plotting function provided by FRASER:
• plotAberrantPerSample()
• plotVolcano()
• plotExpression()
• plotQQ()
• plotExpectedVsObservedPsi()
• plotCountCorHeatmap()
• plotFilterExpression()
• plotFilterVariability()
• plotEncDimSearch()

For a detailed description of each plot function please see the details. Most of the functions share the same parameters.

plotAberrantPerSample: The number of aberrant events per sample are plotted sorted by rank. The ... parameters are passed on to the aberrant function.

plotVolcano: the volcano plot is sample-centric. It plots for a given sample and psi type the negative log10 nominal P-values against the delta psi values for all splice sites or aggregates by gene if requested.

plotExpression: This function plots for a given site the read count at this site (i.e. K) against the total coverage (i.e. N) for the given psi type ($\psi_5$, $\psi_3$, or $\theta$ (SE)) for all samples.

plotQQ: the quantile-quantile plot for a given gene or if global is set to TRUE over the full data set. Here the observed P-values are plotted against the expected ones in the negative log10 space.

plotExpectedVsObservedPsi: A scatter plot of the observed psi against the predicted psi for a given site.

plotCountCorHeatmap: The correlation heatmap of the count data either of the full data set (i.e. sample-sample correlations) or of the top x most variable junctions (i.e. junction-sample correlations). By default the values are log transformed and row centered. The ... arguments are passed to the pheatmap function.

plotFilterExpression: The distribution of FPKM values. If the FraserDataSet object contains the passedfilter column, it will plot both FPKM distributions for the expressed introns and for the filtered introns.

plotFilterVariability: The distribution of maximal delta Psi values. If the FraserDataSet object contains the passedFilter column, it will plot both maximal delta Psi distributions for the variable introns and for the filtered (i.e. non-variable) introns.

plotEncDimSearch: Visualization of the hyperparameter optimization. It plots the encoding dimension against the achieved loss (area under the precision-recall curve). From this plot the optimum should be chosen for the q in fitting process.

Value

If base R graphics are used nothing is returned else the plotly or the gplot object is returned.
psiTypes

Examples

# create full FRASER object
fds <- makeSimulatedFraserDataSet(m=40, j=200)
fds <- calculatePSIValues(fds)
fds <- filterExpressionAndVariability(fds, filter=FALSE)
# this step should be done for all splicing metrics and more dimensions
fds <- optimHyperParams(fds, "psi5", q_param=c(2,5,10,25))
fds <- FRASER(fds)

# QC plotting
plotFilterExpression(fds)
plotFilterVariability(fds)
plotCountCorHeatmap(fds, "theta")
plotCountCorHeatmap(fds, "theta", normalized=TRUE)
plotEncDimSearch(fds, type="psi5")

# extract results
plotAberrantPerSample(fds, aggregate=FALSE)
plotVolcano(fds, "sample1", "psi5")

# dive into gene/sample level results
res <- results(fds)
res
plotExpression(fds, result=res[1])
plotQQ(fds, result=res[1])
plotExpectedVsObservedPsi(fds, type="psi5", res=res[1])

---

### psiTypes

#### Available psi types

<table>
<thead>
<tr>
<th>Description</th>
<th>Available psi types</th>
</tr>
</thead>
<tbody>
<tr>
<td>Available psi types</td>
<td></td>
</tr>
</tbody>
</table>

#### Usage

psiTypes

#### Format

An object of class character of length 3.

#### Examples

# to show available psi types:
psiTypes
results, FraserDataSet-method

Extracting results and aberrant splicing events

Description

The result function extracts the results from the given analysis object based on the given options and cutoffs. The aberrant function extracts aberrant splicing events based on the given cutoffs.

Usage

```r
## S4 method for signature 'FraserDataSet'
results(
  object,
  sampleIDs = samples(object),
  padjCutoff = 0.05,
  zScoreCutoff = NA,
  deltaPsiCutoff = 0.3,
  minCount = 5,
  psiType = c("psi3", "psi5", "theta"),
  additionalColumns = NULL,
  BPPARAM = bpparam(),
  ...
)

resultsByGenes(res, geneColumn = "hgncSymbol", method = "BY")

## S4 method for signature 'FraserDataSet'
aberrant(
  object,
  type = currentType(object),
  padjCutoff = 0.05,
  deltaPsiCutoff = 0.3,
  zScoreCutoff = NA,
  minCount = 5,
  by = c("none", "sample", "feature"),
  aggregate = FALSE,
  ...
)
```

Arguments

- **object** A FraserDataSet object
- **sampleIDs** A vector of sample IDs for which results should be retrieved
- **padjCutoff** The FDR cutoff to be applied or NA if not requested.
- **zScoreCutoff** The z-score cutoff to be applied or NA if not requested.
deltaPsiCutoff  The cutoff on delta psi or NA if not requested.
minCount    The minimum count value of the total coverage of an intron to be considered as significant.
psiType     The psi types for which the results should be retrieved.
additionalColumns  Character vector containing the names of additional columns from mcols(fds) that should appear in the result table (e.g. ensembl_gene_id). Default is NULL, so no additional columns are included.
BPPARAM   The BiocParallel parameter.
res         Result as created with results()
geneColumn  The name of the column in mcols(res) that contains the gene symbols.
method      The p.adjust method that is being used to adjust p values per sample.
type        Splicing type (psi5, psi3 or theta)
by          By default none which means no grouping. But if sample or feature is specified the sum by sample or feature is returned
aggregate   If TRUE the returned object is based on the grouped features

Value
For results: GRanges object containing significant results. For aberrant: Either a of logical values of size introns/genes x samples if "by" is NA or a vector with the number of aberrant events per sample or feature depending on the vaule of "by"

Examples
# get data, fit and compute p-values and z-scores
fds <- createTestFraserDataSet()

# extract results: for this example dataset, z score cutoff of 2 is used to # get at least one result and show the output
res <- results(fds, padjCutoff=NA, zScoreCutoff=3, deltaPsiCutoff=0.05)
res

# aggregate the results by genes (gene symbols need to be annotated first # using annotateRanges() function)
resultsByGenes(res)

# get aberrant events per sample: on the example data, nothing is aberrant # based on the adjusted p-value
aberrant(fds, type="psi5", by="sample")

# get aberrant events per gene (first annotate gene symbols)
fds <- annotateRangesWithTxDb(fds)
aberrant(fds, type="psi5", by="feature", zScoreCutoff=2, padjCutoff=NA,
aggregate=TRUE)

# find aberrant junctions/splice sites
aberrant(fds, type="psi5")

samples Getter/Setter methods for the FraserDataSet

Description

The following methods are getter and setter methods to extract or set certain values of a Fraser-
DataSet object.
samples sets or gets the sample IDs; condition: nonSplicedReads return a RangedSumma-
rizedExperiment object containing the counts for the non spliced reads overlapping splice sites in
the fds.

Usage

samples(object)
samples(object) <- value
condition(object)
condition(object) <- value
bamFile(object)
bamFile(object) <- value
name(object)
name(object) <- value
strandSpecific(object)
strandSpecific(object) <- value
pairedEnd(object)
pairedEnd(object) <- value
workingDir(object)
workingDir(object) <- value
scanBamParam(object)
samples

scanBamParam(object) <- value

nonSplicedReads(object)

nonSplicedReads(object) <- value

## S4 method for signature 'FraserDataSet'
samples(object)

## S4 replacement method for signature 'FraserDataSet'
samples(object) <- value

## S4 method for signature 'FraserDataSet'
condition(object)

## S4 replacement method for signature 'FraserDataSet'
condition(object) <- value

## S4 method for signature 'FraserDataSet'
bamFile(object)

## S4 replacement method for signature 'FraserDataSet'
bamFile(object) <- value

## S4 method for signature 'FraserDataSet'
name(object)

## S4 replacement method for signature 'FraserDataSet'
name(object) <- value

## S4 method for signature 'FraserDataSet'
workingDir(object)

## S4 replacement method for signature 'FraserDataSet'
workingDir(object) <- value

## S4 method for signature 'FraserDataSet'
strandSpecific(object)

## S4 replacement method for signature 'FraserDataSet'
strandSpecific(object) <- value

## S4 method for signature 'FraserDataSet'
pairedEnd(object)

## S4 replacement method for signature 'FraserDataSet'
pairedEnd(object) <- value
## S4 method for signature 'FraserDataSet'
scanBamParam(object)

## S4 replacement method for signature 'FraserDataSet'
scanBamParam(object) <- value

## S4 method for signature 'FraserDataSet'
nonSplicedReads(object)

## S4 replacement method for signature 'FraserDataSet'
nonSplicedReads(object) <- value

FRASER.mcols.get(x, type = NULL, ...)

FRASER.rowRanges.get(x, type = NULL, ...)

mapSeqlevels(fds, style = "UCSC", ...)

### Arguments

- **object**: A FraserDataSet object.
- **value**: The new value that should replace the current one.
- **x**: A FraserDataSet object.
- **type**: The psi type (psi3, psi5 or theta)
- **...**: Further parameters. For mapSeqLevels: further parameters passed to GenomeInfoDb::mapSeqleveles().
- **fds**: FraserDataSet
- **style**: The style of the chromosome names.

### Value

Getter method return the respective current value.

### Author(s)

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### Examples

```r
def <- createTestFraserDataSet()
samples(fds)
samples(fds) <- 1:dim(fds)[2]
condition(fds)
condition(fds) <- 1:dim(fds)[2]
bamFile(fds) # file.paths or objects of class BamFile
bamFile(fds) <- file.path("bamfiles", samples(fds), "rna-seq.bam")```
name(fds) <- "My Analysis"
workingDir(fds) <- tempdir()
strandSpecific(fds) <- TRUE
strandSpecific(fds) <- "reverse"
scanBamParam(fds) <- ScanBamParam(mapqFilter=30)
nonSplicedReads(fds)
rowRanges(fds)
rowRanges(fds, type="theta")
mcols(fds, type="psi5")
mcols(fds, type="theta")
seqlevels(fds)
seqlevels(mapSeqlevels(fds, style="UCSC"))
seqlevels(mapSeqlevels(fds, style="Ensembl"))
seqlevels(mapSeqlevels(fds, style="dbSNP"))

Description

Providing subsetting by indices through the single-bracket operator

Usage

## S3 method for class 'FRASER'
subset(x, i, j, by = c("j", "ss"), ..., drop = FALSE)

## S4 method for signature 'FraserDataSet,ANY,ANY,ANY'
x[i, j, by = c("j", "ss"), ..., drop = FALSE]

Arguments

x
- A FraserDataSet object
i
- A integer vector to subset the rows/ranges
j
- A integer vector to subset the columns/samples
by
- a character (j or ss) defining if we subset by junctions or splice sites
... Parameters currently not used or passed on
drop
- No dimension reduction is done. And the drop parameter is currently not used at all.
Value

A subsetted FraserDataSet object

Examples

```r
fds <- createTestFraserDataSet()
fds[1:10, 2:3]
fds[, samples(fds) %in% c("sample1", "sample2")]
fds[1:10, by = "ss"]
```
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